10

What is claimed is:

- Isolated Apo-2 polypeptide having at least 80% amino acid sequence identity with native sequence Apo-2 polypeptide comprising amino acid residues 1 to 411 of SEQ ID NO:1.
- 5 2. The Apo-2 polypeptide of claim 1 wherein said polypeptide has at least 90% amino acid sequence identity.
 - The Apo-2 polypeptide of claim 2 wherein said polypeptide has at least 95% amino acid sequence identity.
 - Isolated Apo-2 polypeptide comprising amino acid residues 1 to 411 of SEQ ID NO:1.
 - Isolated extracellular domain sequence of Apo-2 polypeptide comprising amino acid residues 54 to 182 of SEQ ID NO:1.
 - The extracellular domain sequence of claim 5 comprising amino acid residues 1 to 182 of SEQ ID NO:1.
 - Isolated death domain sequence of Apo-2 polypeptide comprising amino acid residues 324 to 391 of SEQ ID NO:1.
 - 8. A chimeric molecule comprising the Apo-2 polypeptide of claim 1 or the extracellular domain sequence of claim 5 fused to a heterologous amino acid sequence.
 - The chimeric molecule of claim 8 wherein said heterologous amino acid sequence is an epitope tag sequence.
 - 10. The chimeric molecule of claim 8 wherein said heterologous amino acid sequence is an immunoglobulin sequence.
 - The chimeric molecule of claim 10 wherein said immunoglobulin sequence is an IgG.
 - 12. Isolated nucleic acid comprising a DNA encoding the polypeptide of claim 1, the extracellular domain sequence of claim 5, or the death domain sequence of claim 7.
- 13. The nucleic acid of claim 12 wherein said DNA encodes an Apo-2
 polypeptide comprising amino acid residues 1 to 411 of SEQ ID
 NO:1.
 - 14. A vector comprising the nucleic acid of claim 12.
 - 15. The vector of claim 14 operably linked to control sequences recognized by a host cell transformed with the vector.
- 35 16. The vector of claim 14 comprising ATCC deposit accession number 209021.
 - 17. A host cell comprising the vector of claim 14.

5

10

- 18. The host cell of claim 17 comprising a CHO cell.
- 19. The host cell of claim 17 comprising E. coli.
- 20. The host cell of claim 17 comprising a yeast cell.
- 21. A process of producing an Apo-2 polypeptide comprising culturing the host cell of claim 17 under conditions sufficient to express Apo-2 polypeptide and recovering the expressed Apo-2 polypeptide from the culture.
 - 22. An Apo-2 polypeptide which is obtained or obtainable by expressing the polypeptide encoded by the cDNA insert in ATCC deposit accession number 209021.
 - 23. A non-human, transgenic animal which contains cells that express DNA encoding Apo-2 polypeptide.
 - 24. The animal of claim 23 which is a mouse or rat.
 - 25. A non-human, knockout animal which contains cells having an altered gene encoding Apo-2 polypeptide.
 - 26. The animal of claim 25 which is a mouse or rat.
 - 27. An antibody which specifically binds to an Apo-2 polypeptide.
 - 28. The antibody of claim 27 which is a monoclonal antibody.
 - 29. The antibody of claim 27 comprising an agonistic antibody.
- 30. The antibody of claim 27 comprising a blocking antibody.
- 31. The antibody of claim 24 comprising a chimeric antibody.
- The antibody of claim 28 wherein said antibody is an IgG antibody.
- 33. The antibody of claim 28 wherein said antibody comprises an Fab fragment.
- 34. The antibody of claim 28 wherein said antibody comprises a scFv fragment.
- 35. The antibody of claim 28 wherein said antibody comprises a F(ab')2 fragment.
- 30 36. The antibody of claim 27 wherein said antibody comprises a human antibody.
 - 37. The antibody of claim 28 having the biological characteristics of the monoclonal antibody produced by the hybridoma cell line deposited as ATCC accession number HB-12456.
- 35 38. The antibody of claim 28 wherein the antibody binds to the same epitope as the epitope to which the monoclonal antibody

30

10

- produced by the hybridoma cell line deposited as ATCC accession number HB-12456 binds.
- A hybridoma cell line which produces the antibody of claim 28.
- 5 40. The hybridoma cell line deposited as ATCC accession number HB-12456.
 - The monoclonal antibody produced by the hybridoma cell line deposited as ATCC accession number HB-12456.
 - The antibody of claim 27 wherein said antibody is a singlechain antibody.
 - 43. The antibody of claim 42 wherein said antibody comprises the 16E2 antibody.
 - 44. The antibody of claim 42 wherein said antibody comprises the 20E6 antibody.
 - 45. The antibody of claim 42 wherein said antibody comprises the 24C4 antibody.
 - 46. The antibody of claim 42 wherein said antibody is fused to an epitope tag sequence.
 - 47. A chimeric molecule comprising the antibody of claim 27 fused to a heterologous amino acid sequence.
 - 48. The chimeric molecule of claim 47 wherein said heterologous amino acid sequence comprises an immunoglobulin sequence.
 - 49. A dimeric molecule comprising the Apo-2 antibody of claim 27 and a heterologous antibody.
- 25 50. A homodimeric molecule comprising a first Apo-2 antibody and a second Apo-2 antibody.
 - Isolated nucleic acid comprising DNA encoding the Apo-2 antibody of claim 43.
 - 52. Isolated nucleic acid comprising DNA encoding the antibody of claim 44.
 - Isolated nucleic acid comprising DNA encoding the antibody of claim 45.
 - 54. A vector comprising the nucleic acid of claim 51, 52, or 53.
 - 55. A host cell comprising the vector of claim 54.
- 35 56. A method of producing an Apo-2 antibody comprising culturing the host cell of claim 55 under conditions wherein the DNA is expressed.

- 57. A composition comprising the antibody of claim 27 and a carrier.
- 58. The composition of claim 57 wherein said carrier is a pharmaceutically-acceptable carrier.
- 5 59. A method of inducing apoptosis in mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of the Apo-2 agonistic antibody of claim 29.
 - The method of claim 59 wherein said agonistic antibody comprises a single-chain antibody.
- 10 61. A method of treating mammalian cancer cells comprising exposing mammalian cancer cells to an agent which activates Apo-2.
 - 62. The method of claim 61 wherein said agent comprises an agonistic Apo-2 antibody.
 - 63. An article of manufacture comprising a container and a composition contained within said container, wherein the composition includes Apo-2 polypeptide or Apo-2 antibody.
 - 64. The article of manufacture of claim 63 further comprising instructions for using the Apo-2 polypeptide or Apo-2 antibody in vivo or ex vivo.